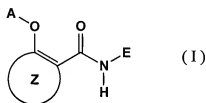


Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1. (Currently Amended): A ~~medicament~~ method for preventive and/or therapeutic treatment of ~~Alzheimer's disease~~ neural diseases caused by activation of NF- κ B and AP-1 in a mammal, which comprises ~~as an active ingredient~~ administering to a mammal a preventively and/or therapeutically effective amount of a substance selected from the group consisting of a compound represented by the following general formula (I), and a pharmacologically acceptable salt thereof, and a hydrate thereof, and a solvate thereof:



wherein A represents hydrogen atom or acetyl group,

E represents a 2,5-di-substituted or phenyl group wherein at least one of said substituents is trifluoromethyl group.

a 3,5-di-substituted phenyl group wherein at least one of said substituents is trifluoromethyl group, or a monocyclic or a fused polycyclic heteroaryl group which may be substituted, provided that the compound wherein said heteroaryl group is (1) a fused polycyclic heteroaryl group wherein the ring which binds directly to CONH group in the formula (I) is a benzene ring, (2) unsubstituted thiazol-2-yl group, or (3) unsubstituted benzothiazol-2-yl group is excluded,

a 2-thiazolyl group which is substituted with one or more substituents selected from the group consisting of

a halogen atom,

an alkyl group which may be substituted with one or more substituents selected from the group consisting of

a carboxy group and

an alkoxy-carbonyl group,

a halogenated alkyl group,

a cyano group,

an aryl group which may be substituted with one or more substituents selected from the group consisting of

a halogen atom,

a halogenated alkyl group and

an alkoxy group,

an alkyl-carbonyl group,

an alkoxy-carbonyl group,

a monocyclic non-aromatic heterocyclic group which may be substituted with one or more substituents selected from the group consisting of

an alkyl group and

an aryl group,

an aralkyl group,

an aryl-carbonyl group,

a carbamoyl group which may be substituted with one or more substituents selected from the group consisting of

an alkyl group and

an aralkyl group, and

a carboxy group,

ring Z represents ~~an arene~~ a benzene ring which may have one or more substituents selected from the group consisting of

a halogen atom,

a nitro group,

a cyano group,

a hydroxy group,

an alkoxy group,

an alkyl group which may be substituted with one or more substituents selected from the group consisting of

a hydroxy group,

an aralkyl-oxy-imino group and

an alkoxy-imino group,

an alkenyl group which may be substituted with one or more substituents selected from the group consisting of

an aryl group,

a cyano group,

an alkoxy-carbonyl group and

a carboxy group,

an alkynyl group which may be substituted with one or more substituents selected from the group consisting of

an aryl group and

a tri(alkyl)silyl group,

a halogenated alkyl group,

an aryl group which may be substituted with one or more substituents selected from the group consisting of

a halogen atom and

a halogenated alkyl group,

an aralkyl group,

a monocyclic or a fused polycyclic heteroaryl group which may be substituted with one or more alkyl groups,

an alkyl-carbonyl group,

a monocyclic non-aromatic heterocyclic-carbonyl group which may be substituted with one or more aralkyl groups,

a monocyclic heteroaryl-sulfonyl group,

a carboxy group,

an alkoxy-carbonyl group,

a carbamoyl group which may be substituted with one or more substituents selected from the group consisting of

an aryl group which may be substituted with one or more halogenated alkyl groups and

an alkyl group,

a sulfamoyl group which may be substituted with one or more substituents selected from the group consisting of

an aryl group which may be substituted with one or more halogenated alkyl groups and

an alkyl group,

an amino group which may be substituted with one or more substituents selected from the group consisting of

an alkyl group,

an alkyl-carbonyl group,

an aryl-carbonyl group,

an alkyl-sulfonyl group and

an aryl-sulfonyl group,

an ureido group which may be substituted with one or more aryl groups,

a thioureido group which may be substituted with one or more aryl groups, and

a diazenyl group which may be substituted with one or more aryl groups wherein the aryl groups may be substituted with one or more substituents selected from the group consisting of

a nitro group and

a monocyclic heteroaryl-sulfamoyl group,

in addition to the group represented by formula -O-A wherein A has the same meaning as that defined above and the group represented by formula -CONH-E wherein E has the same meaning as that defined above, ~~or a heteroarene which may have one or more substituents in addition to the group represented by formula -O-A wherein A has the~~

~~same meaning as that defined above and the group represented by formula—CONH—E~~
~~wherein E has the same meaning as that defined above.~~

2-12. (Canceled)

13. (New) The method according to claim 1, wherein the mammal is a human.

14. (New) The method according to claim 1, wherein the neural disease is
Alzheimer's disease or epilepsy.

15. (New) The method according to claim 14, wherein the neural disease is
Alzheimer's disease.

16. (New) The method according to claim 14, wherein the neural disease is
epilepsy.

17. (New) The method according to claim 1, wherein E is a 2,5-di-substituted
phenyl group wherein at least one of said substituents is trifluoromethyl group.

18. (New) The method according to claim 17, wherein E is a 2,5-di-substituted
phenyl group wherein at least one of said substituents is trifluoromethyl group, and the
other substituent is selected from the group consisting of
a halogen atom,

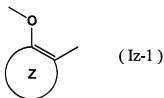
a halogenated alkyl group,
a nitro group,
an alkyl group,
an alkoxy group,
an alkyl-sulfanyl group,
a monocyclic non-aromatic heterocyclic group which may be substituted with one or more halogenated alkyl groups,

an aryl-oxy group which may be substituted with one or more substituents selected from the group consisting of

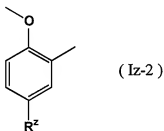
a halogen atom,
an alkoxy group,
an alkyl group and
a cyano group, and
a halogenated alkoxy group.

19. (New) The method according to claim 18, wherein E is a 2-chloro-5-(trifluoromethyl)phenyl group, a 2,5-bis(trifluoromethyl)phenyl group, a 2-fluoro-5-(trifluoromethyl)phenyl group, a 2-nitro-5-(trifluoromethyl)phenyl group, a 2-methyl-5-(trifluoromethyl)phenyl group, a 2-methoxy-5-(trifluoromethyl)phenyl group, a 2-methylsulfanyl-5-(trifluoromethyl)phenyl group, a 2-(1-pyrrolidinyl)-5-(trifluoromethyl)phenyl group, a 2-morpholino-5-(trifluoromethyl)phenyl group, a 2-bromo-5-(trifluoromethyl)phenyl group, a 2-(2-naphthyloxy)-5-(trifluoromethyl)phenyl group, a 2-(2,4-dichlorophenoxy)-5-(trifluoromethyl)phenyl group, a 2-[4-

(trifluoromethyl)piperidin-1-yl]-5-(trifluoromethyl)phenyl group, a 2-(2,2,2-trifluoroethoxy)-5-(trifluoromethyl)phenyl group, a 2-(2-methoxyphenoxy)-5-(trifluoromethyl)phenyl group, a 2-(4-chloro-3,5-dimethylphenoxy)-5-(trifluoromethyl)phenyl group, a 2-piperidino-5-(trifluoromethyl)phenyl group, a 2-(4-methylphenoxy)-5-(trifluoromethyl)phenyl group, a 2-(4-chlorophenoxy)-5-(trifluoromethyl)phenyl group, a 2-(4-cyanophenoxy)-5-(trifluoromethyl)phenyl group or a 2-(4-methoxyphenoxy)-5-(trifluoromethyl)phenyl group, the following partial formula (Iz-1) in the general formula containing ring Z



is represented by the following formula (Iz-2):



wherein R^Z represents a hydrogen atom, a halogen atom, a nitro group, a cyano group, a methoxy group, a methyl group, an isopropyl group, a tert-butyl group, a 1,1,3,3-tetramethylbutyl group, a 2-phenylethen-1-yl group, a 2,2-dicyanoethen-1-yl group, a 2-cyano-2-(methoxycarbonyl)ethen-1-yl group, a 2-carboxy-2-cyanoethen-1-yl group, an ethynyl group, a phenylethynyl group, a (trimethylsilyl)ethynyl group, a trifluoromethyl group, a pentafluoroethyl group, a phenyl group, a 4-(trifluoromethyl)phenyl group, a 4-fluorophenyl group, a 2,4-difluorophenyl group, a 2-phenethyl group, a 1-hydroxyethyl

group, a 1-(methoxyimino)ethyl group, a 1-[(benzyloxy)imino]ethyl group, a 2-thienyl group, a 3-thienyl group, a 1-pyrrolyl group, a 2-methylthiazol-4-yl group, an imidazo[1,2-a]pyridin-2-yl group, a 2-pyridyl group, an acetyl group, an isobutyryl group, a piperidinocarbonyl group, a 4-benzylpiperidinocarbonyl group, a (pyrrol-1-yl)sulfonyl group, a carboxy group, a methoxycarbonyl group, an N-[3,5-bis(trifluoromethyl)phenyl]carbamoyl group, an N,N-dimethylcarbamoyl group, a sulfamoyl group, an N-[3,5-bis(trifluoromethyl)phenyl]sulfamoyl group, an N,N-dimethylsulfamoyl group, an amino group, an N,N-dimethylamino group, an acetylamino group, a benzoylamino group, a methanesulfonylamino group, a benzenesulfonylamino group, a 3-phenylureido group, a (3-phenyl)thioureido group, a (4-nitrophenyl)diazenyl group or a {[4-(pyridin-2-yl)sulfamoyl]phenyl}diazenyl group.

20. (New) The method according to claim 19, wherein A is a hydrogen atom, R^z is a halogen atom.

21. (New) The method according to claim 20, wherein E is a 2,5-bis(trifluoromethyl)phenyl group.

22. (New) The method according to claim 21, wherein R^z is a bromine atom.

23. (New) The method according to claim 17, wherein E is a 2,5-bis(trifluoromethyl)phenyl group.

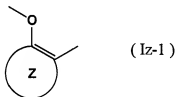
24. (New) The method according to claim 1, wherein E is a 3,5-di-substituted phenyl group wherein at least one of said substituents is trifluoromethyl group.

25. (New) The method according to claim 24, wherein E is a 3,5-di-substituted phenyl group wherein at least one of said substituents is trifluoromethyl group, and the other substituent is selected from the group consisting of

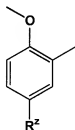
- a halogenated alkyl group,
- a halogen atom,
- an alkoxy group,
- an alkoxy-carbonyl group and
- a carboxy group.

26. (New) The method according to claim 25, wherein E is a 3,5-bis(trifluoromethyl)phenyl group, a 3-fluoro-5-(trifluoromethyl)phenyl group, a 3-bromo-5-(trifluoromethyl)phenyl group, a 3-methoxy-5-(trifluoromethyl)phenyl group, a 3-methoxycarbonyl-5-(trifluoromethyl)phenyl group or a 3-carboxy-5-(trifluoromethyl)phenyl group,

the following partial formula (Iz-1) in the general formula containing ring Z



is represented by the following formula (Iz-2):



(Iz-2)

wherein R^z represents a hydrogen atom, a halogen atom, a nitro group, a cyano group, a methoxy group, a methyl group, an isopropyl group, a tert-butyl group, a 1,1,3,3-tetramethylbutyl group, a 2-phenylethen-1-yl group, a 2,2-dicyanoethen-1-yl group, a 2-cyano-2-(methoxycarbonyl)ethen-1-yl group, a 2-carboxy-2-cyanoethen-1-yl group, an ethynyl group, a phenylethynyl group, a (trimethylsilyl)ethynyl group, a trifluoromethyl group, a pentafluoroethyl group, a phenyl group, a 4-(trifluoromethyl)phenyl group, a 4-fluorophenyl group, a 2,4-difluorophenyl group, a 2-phenethyl group, a 1-hydroxyethyl group, a 1-(methoxyimino)ethyl group, a 1-[(benzyloxy)imino]ethyl group, a 2-thienyl group, a 3-thienyl group, a 1-pyrrolyl group, a 2-methylthiazol-4-yl group, an imidazo[1,2-a]pyridin-2-yl group, a 2-pyridyl group, an acetyl group, an isobutyryl group, a piperidinocarbonyl group, a 4-benzylpiperidinocarbonyl group, a (pyrrol-1-yl)sulfonyl group, a carboxy group, a methoxycarbonyl group, an N-[3,5-bis(trifluoromethyl)phenyl]carbamoyl group, an N,N-dimethylcarbamoyl group, a sulfamoyl group, an N-[3,5-bis(trifluoromethyl)phenyl]sulfamoyl group, an N,N-dimethylsulfamoyl group, an amino group, an N,N-dimethylamino group, an acetylamino group, a benzoylamino group, a methanesulfonylamino group, a benzenesulfonylamino group, a 3-phenylureido group, a (3-phenyl)thioureido group, a (4-nitrophenyl)diazanyl group or a {[4-(pyridin-2-yl)sulfamoyl]phenyl}diazanyl group.

27. (New) The method according to claim 26, wherein A is a hydrogen atom, R² is a halogen atom.

28. (New) The method according to claim 27, wherein E is a 3,5-bis(trifluoromethyl)phenyl group.

29. (New) The method according to claim 28, wherein R² is a chlorine atom.

30. (New) The method according to claim 24, wherein E is a 3,5-bis(trifluoromethyl)phenyl group.

31. (New) The method according to claim 1, wherein E is a 2-thiazolyl group which is substituted with one or more substituents selected from the group consisting of

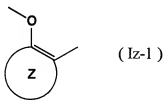
- a halogen atom,
- an alkyl group which may be substituted with one or more substituents selected from the group consisting of
 - a carboxy group and
 - an alkoxy-carbonyl group,
- a halogenated alkyl group,
- a cyano group,
- an aryl group which may be substituted with one or more substituents selected from the group consisting of
 - a halogen atom,

a halogenated alkyl group and
an alkoxy group,
an alkyl-carbonyl group,
an alkoxy-carbonyl group,
a monocyclic non-aromatic heterocyclic group which may be substituted with one or more substituents selected from the group consisting of
an alkyl group and
an aryl group,
an aralkyl group,
an aryl-carbonyl group,
a carbamoyl group which may be substituted with one or more substituents selected from the group consisting of
an alkyl group and
an aralkyl group, and
a carboxy group.

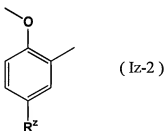
32. (New) The method according to claim 31, wherein E is a 5-bromo-4-[(1,1-dimethyl)ethyl]thiazol-2-yl group, a 5-bromo-4-(trifluoromethyl)thiazol-2-yl group, a 5-cyano-4-[(1,1-dimethyl)ethyl]thiazol-2-yl group, a 4,5-dimethylthiazol-2-yl group, a 5-methyl-4-phenylthiazol-2-yl group, a 5-(4-fluorophenyl)-4-methylthiazol-2-yl group, a 4-methyl-5-[3-(trifluoromethyl)phenyl]thiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-ethylthiazol-2-yl group, a 4-ethyl-5-phenylthiazol-2-yl group, a 4-isopropyl-5-phenylthiazol-2-yl group, a 4-butyl-5-phenylthiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-

5-[(2,2-dimethyl)propionyl]thiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-(ethoxycarbonyl)thiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-piperidinethiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-morpholinethiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-(4-methylpiperazin-1-yl)thiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]-5-(4-phenylpiperazin-1-yl)thiazol-2-yl group, a 5-carboxymethyl-4-phenylthiazol-2-yl group, a 4,5-diphenylthiazol-2-yl group, a 4-benzyl-5-phenylthiazol-2-yl group, a 5-phenyl-4-(trifluoromethyl)thiazol-2-yl group, a 5-acetyl-4-phenylthiazol-2-yl group, a 5-benzoyl-4-phenylthiazol-2-yl group, a 5-ethoxycarbonyl-4-phenylthiazol-2-yl group, a 5-ethoxycarbonyl-4-(pentafluorophenyl)thiazol-2-yl group, a 5-methylcarbamoyl-4-phenylthiazol-2-yl group, a 5-ethylcarbamoyl-4-phenylthiazol-2-yl group, a 5-isopropylcarbamoyl-4-phenylthiazol-2-yl group, a 5-(2-phenylethyl)carbamoyl-4-phenylthiazol-2-yl group, a 5-ethoxycarbonyl-4-(trifluoromethyl)thiazol-2-yl group, a 5-carboxy-4-[(1,1-dimethyl)ethyl]thiazol-2-yl group, a 5-(ethoxycarbonyl)methyl-4-phenylthiazol-2-yl group, a 5-carboxy-4-phenylthiazol-2-yl group, a 5-propylcarbamoyl-4-phenylthiazol-2-yl group, a 5-methylthiazol-2-yl group, a 4-[(1,1-dimethyl)ethyl]thiazol-2-yl group, a 4-phenylthiazol-2-yl group, a 4-[3,5-bis(trifluoromethyl)phenyl]thiazol-2-yl group, a 4-(2,4-dichlorophenyl)thiazol-2-yl group, a 4-(3,4-dichlorophenyl)thiazol-2-yl group, a 4-[4-(trifluoromethyl)phenyl]thiazol-2-yl group, a 4-(2,5-difluorophenyl)thiazol-2-yl group, a 4-(4-methoxyphenyl)thiazol-2-yl group, a 4-[3-(trifluoromethyl)phenyl]thiazol-2-yl group or a 4-(pentafluorophenyl)thiazol-2-yl group,

the following partial formula (Iz-1) in the general formula containing ring Z



is represented by the following formula (Iz-2):



wherein R^z represents a hydrogen atom, a halogen atom, a nitro group, a cyano group, a methoxy group, a methyl group, an isopropyl group, a tert-butyl group, a 1,1,3,3-tetramethylbutyl group, a 2-phenylethen-1-yl group, a 2,2-dicyanoethen-1-yl group, a 2-cyano-2-(methoxycarbonyl)ethen-1-yl group, a 2-carboxy-2-cyanoethen-1-yl group, an ethynyl group, a phenylethynyl group, a (trimethylsilyl)ethynyl group, a trifluoromethyl group, a pentafluoroethyl group, a phenyl group, a 4-(trifluoromethyl)phenyl group, a 4-fluorophenyl group, a 2,4-difluorophenyl group, a 2-phenethyl group, a 1-hydroxyethyl group, a 1-(methoxyimino)ethyl group, a 1-[(benzyloxy)imino]ethyl group, a 2-thienyl group, a 3-thienyl group, a 1-pyrrolyl group, a 2-methylthiazol-4-yl group, an imidazo[1,2-a]pyridin-2-yl group, a 2-pyridyl group, an acetyl group, an isobutyryl group, a piperidinocarbonyl group, a 4-benzylpiperidinocarbonyl group, a (pyrrol-1-yl)sulfonyl group, a carboxy group, a methoxycarbonyl group, an N-[3,5-bis(trifluoromethyl)phenyl]carbamoyl group, an N,N-dimethylcarbamoyl group, a sulfamoyl group, an N-[3,5-bis(trifluoromethyl)phenyl]sulfamoyl group, an N,N-dimethylsulfamoyl group, an amino group, an N,N-dimethylamino group, an acetylamino

group, a benzoylamino group, a methanesulfonylamino group, a benzenesulfonylamino group, a 3-phenylureido group, a (3-phenyl)thioureido group, a (4-nitrophenyl)diazenyl group or a {[4-(pyridin-2-yl)sulfamoyl]phenyl}diazenyl group.

33. (New) The method according to claim 32, wherein A is a hydrogen atom, R^z is a halogen atom.

34. (New) The method according to claim 33, wherein E is a 4-[(1,1-dimethyl)ethyl]-5-[(2,2-dimethyl)propionyl]thiazol-2-yl group.